(FILE 'HOME' ENTERED AT 10:31:44 ON 01 FEB 2011)

FILE 'MEDLINE, CAPLUS, BIOSIS, SCISEARCH, LIFESCI' ENTERED AT 10:31:59 ON 01 FEB 2011

- L1 228783 S (PLURIPOTENT OR EMBRYONIC) (4A) CELL
- L2 8534 S EMBRYOID(W) (BODY OR BODIES)
- L3 6715 S L1(P)L2
- L4 2452 S L1(9A)L2
- L5 1181 DUP REM L4 (1271 DUPLICATES REMOVED)
- L6 1171 S EMBRYONIC (W) GERM (W) CELL
- L7 67 S L6(P)L2
- L8 25 DUP REM L7 (42 DUPLICATES REMOVED)
- => d au ti so pi 1-25 18
- L8 ANSWER 1 OF 25 MEDLINE on STN DUPLICATE 1
- AU Hiller Marc; Liu Cyndi; Blumenthal Paul D; Gearhart John D; Kerr Candace L
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- SO Stem cells and development, (2011 Feb) Vol. 20, No. 2, pp. 351-61. Electronic Publication: 2010-10-21. Journal code: 101197107. E-ISSN: 1557-8534. L-ISSN: 1547-3287.
- L8 ANSWER 2 OF 25 CAPLUS COPYRIGHT 2011 ACS on STN
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- TI Ascorbic acid induces differentiation of human embryonic germ cells towards cardiomyocytes
- SO Jiangsu Yiyao (2010), 36(13), 1551-1554, C2 CODEN: CIYADX; ISSN: 0253-3685
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- SO Stem cells and development, (2010 Feb) Vol. 19, No. 2, pp. 195-202. Journal code: 101197107. E-ISSN: 1557-8534. L-ISSN: 1547-3287.
- L8 ANSWER 4 OF 25 MEDLINE on STN DUPLICATE 3
- AU Hillel Alexander T; Varghese Shyni; Petsche Jennifer; Shamblott Michael J; Elisseeff Jennifer H
- TI Embryonic germ cells are capable of adipogenic differentiation in vitro and in vivo.
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- L8 ANSWER 5 OF 25 SCISEARCH COPYRIGHT (c) 2011 The Thomson Corporation on STN
- AU Crane, Janet L.; Hsu, Stephanie; Germain-Lee, Emily L. (Reprint); Shamblott, Michael J.; Shamblott, Michael J.; Axelman, Joyce; Levine, Michael A.; Levine, Michael A.
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- AU Hua Jinlian; Yu Haisheng; Liu Sheng; Dou Zhongying; Sun Yadong; Jing Xiaoqi; Yang Chunrong; Lei Anmin; Wang Huayan; Gao Zhimin
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- L8 ANSWER 7 OF 25 MEDLINE on STN DUPLICATE 5
- AU Wang Juan; Jiao Fei; Pan Xiao-Hong; Xie Shu-Yang; Li Zun-Ling; Niu Xin-Hua; Du Li-Xin
- TI Directed differentiation of chick embryonic germ cells into neural cells using retinoic acid induction in vitro.
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- L8 ANSWER 8 OF 25 CAPLUS COPYRIGHT 2011 ACS on STN DUPLICATE 6
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- SO Dongwu Xuebao (2008), 54(5), 855-860 CODEN: TWHPA3; ISSN: 0001-7302
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- L8 ANSWER 11 OF 25 BIOSIS COPYRIGHT (c) 2011 The Thomson Corporation on STN
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- TI Musculoskeletal differentiation of cells derived from human embryonic germ cells
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- TI De-differentiation and re-differentiation of somatic cells and production of cells for cell therapies
- SO PCT Int. Appl. CODEN: PIXXD2

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- L8 ANSWER 20 OF 25 BIOSIS COPYRIGHT (c) 2011 The Thomson Corporation on STN
- AU Kerr, D. A. [Reprint Author]; Llado, J. [Reprint Author]; Shamblott, M. J. [Reprint Author]; Maragakis, N. J. [Reprint Author]; Irani, D. N. [Reprint Author]; Crawford, T. O. [Reprint Author]; Gearhart, J. D. [Reprint Author]; Rothstein, J. D. [Reprint Author]
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- L8 ANSWER 21 OF 25 CAPLUS COPYRIGHT 2011 ACS on STN
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## => d ab 20-25 18

- L8 ANSWER 20 OF 25 BIOSIS COPYRIGHT (c) 2011 The Thomson Corporation on  ${\tt STN}$
- We have investigated the potential of human pluripotent cells to restore AΒ function in rats paralyzed with a virus-induced motor neuronopathy. Cells derived from embryonic germ cells introduced into the cerebrospinal fluid were distributed extensively over the rostral-caudal length of the spinal cord and migrated into the spinal cord parenchyma. Some of the transplanted human cells expressed the neuroglial progenitor marker nestin, while others expressed immunohistochemical markers characteristic of astrocytes or mature neurons. Rare transplanted cells developed immunoreactivity to choline acetyl-transferase and sent axons into the sciatic nerve as detected by retrograde labeling. Paralyzed animals transplanted with embryoid body -derived cells partially recovered motor function 12 and 24 weeks post transplantation, whereas control animals remained paralyzed. Transplanted embryoid body-derived cells protected host neurons from death and facilitated reafferentation of motor neuron cell bodies. We conclude that cells derived from human pluripotent stem cells have the capacity to restore neurologic function in animals with diffuse motor neuron disease and may present a potential strategy for treating individuals with amyotrophic lateral sclerosis or spinal muscular atrophy.
- L8 ANSWER 21 OF 25 CAPLUS COPYRIGHT 2011 ACS on STN
- EG4 cells derived from primordial germ cells (PGCs) of 10.5 d post coitum AΒ 129/svJ mouse embryos can be used as a model system for in vitro differentiation study due to their pluripotential development ability. EG4 cell lines with stable expression of kinase-neg. EGFR cDNA, designated EG4-EGFRd, were generated by gene transfection. We found that: (i) EG4-EGFRd cells share the similar morphol. and growing character with wildtype cells that can maintain undifferentiated state in long term culture. (ii) Treatment of EG4 cells with RA resulted in differentiation of adipocyte, while in mutant clones of EG4-EGFRd, adipocytes were sparse or absent under the same condition, indicating the role of EGFR expressed during adipocyte development. (iii) Histol. anal. showed that predominant tissues in teratocarcinomas derived from EG4-EGFRd cells and wildtype cells are different. A large amount of undifferentiated cells was present in those coming from mutant cell clones. In addition some cardiac and skeletal muscles are prominently differentiated cell types. EG4 wildtype cells produced multiple differentiated cell types of three primary germ layers such as cartilage, epithelia and neural tube. These studies suggested that EGFR-dependent differentiation was inhibited in kinase-neg. EG4 cells.
- L8 ANSWER 22 OF 25 MEDLINE on STN DUPLICATE 15
- AB Human pluripotent stem cells (hPSCs) have been derived from the inner cell mass cells of blastocysts (embryonic stem cells) and primordial germ cells of the developing gonadal ridge (embryonic germ

cells). Like their mouse counterparts, hPSCs can be maintained in culture in an undifferentiated state and, upon differentiation, generate a wide variety of cell types. Embryoid body (EB) formation is a requisite step in the process of in vitro differentiation of these stem cells and has been used to derive neurons and glia, vascular endothelium, hematopoietic cells, cardiomyocytes, and glucose-responsive insulin-producing cells from mouse PSCs. EBs generated from human embryonic germ cell cultures have also been found to contain a wide variety of cell types, including neural cells, vascular endothelium, muscle cells, and endodermal derivatives. Here, we report the isolation and culture of cells from human EBs as well as a characterization of their gene expression during growth in several different culture environments. These heterogeneous cell cultures are capable of robust and long-term [>70 population doublings (PD)] proliferation in culture, have normal karyotypes, and can be cryopreserved, clonally isolated, and stably transfected. Cell cultures and clonal lines retain a broad pattern of gene expression including simultaneous expression of markers normally associated with cells of neural, vascular/hematopoietic, muscle, and endoderm lineages. The growth and expression characteristics of these EB-derived cells suggest that they are relatively uncommitted precursor or progenitor cells. EB-derived cells may be suited to studies of human cell differentiation and may play a role in future transplantation therapies.

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At present embryonic stem (ES) cells with confirmed pluripotential AB properties are only available in the mouse. Recently, we were able to isolate, culture and genetically transform primordial germ cell (PGC)-derived cells from pig embryos and demonstrate their ability to contribute to chimera development in the pig. In order to determine whether the system we developed could be used to isolate embryonic germ (EG) cells from other mammalian species, we placed isolated PGCs from cattle, goats, rabbits and rats in culture. Briefly, PGCs were isolated from fetuses of cow (day 30-50), goat (day 25), rabbit (day 15-18) and rat (day 11-12), and plated on. STO feeder cells in Dulbecco's modified Eagle's medium (DMEM): Ham's F10 medium (1:1) supplemented with 0.01 mM nonessential amino acids, 2 mM L-glutamine, 0.1 mM beta- mercaptoethnol, soluble recombinant human stem cell factor (SCF; 40ng/ml), human basic fibroblast growth factor (bFGF; 20ng/ml) and human leukemia inhibitory factor (LIF; 20ng/ml). For maintenance of the cells, colonies were passed to fresh feeders every 7-10 days. In all species tested, we were able to obtain and maintain colonies with ES-like morphology. Their developmental potential was tested by alkaline phosphatase (AP) staining and in vitro differentiation assay. For genetic transformation, cells were electroporated with a construct containing the green fluorescent protein (GFP) under the control of the cytomegalovirus (CMV) promoter. (GFP-expressing colonies were detected in cattle, rabbits and rats. These results suggest that PGC-derived cells from cattle, goats, rabbits and rats can be isolated, cultured, and genetically transformed, and provide the basis for analyzing their developmental potential and their possible use for the precise genetic modification of these species.

ANSWER 24 OF 25 CAPLUS COPYRIGHT 2011 ACS on STN

Induction of hematopoiesis in an embryonic germ (EG) cell line derived from mouse primordial germ cells (PGCs) was examined When single undifferentiated EG-1 cells were inoculated directly into the methylcellulose medium, both primitive and definitive erythropoiesis were seen in embryoid bodies derived from the EG cells as observed in ES cells, and production of myeloid cell lineages was stimulated by IL-3. These results indicate that EG cells acquired in vitro potency to differentiate toward

hematopoietic cells, although they were derived from PGC and are distinct from inner cell mass-derived ES cells with regard to gene expression and patterns of DNA methylation corresponding to genomic imprinting. It turns out that they are useful for study of cell differentiation in the animals whose ES cells are not available.

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AB

Human pluripotent stem cells would be invaluable for in vitro studies of aspects of human embryogenesis. With the goal of establishing pluripotent stem cell lines, gonadal ridges and mesenteries containing primordial germ cells (PGCs, 5-9 weeks postfertilization) were cultured on mouse STO fibroblast feeder layers in the presence of human recombinant leukemia inhibitory factor, human recombinant basic fibroblast growth factor, and forskolin. Initially, single PGCs in culture were visualized by alkaline phosphatase activity staining. Over a period of 7-21 days, PGCs gave rise to large multicellular colonies resembling those of mouse pluripotent stem cells termed embryonic stem and embryonic germ (EG) cells. Throughout the culture period most cells within the colonies continued to be alkaline phosphatase-positive and tested positive against a panel of five immunological markers (SSEA-1, SSEA-3, SSEA-4, TRA-1-60, and TRA-1-81) that have been used routinely to characterize embryonic stem and EG cells. The cultured cells have been continuously passaged and found to be karyotypically normal and stable. Both XX and XY cell cultures have been obtained. Immunohistochemical analysis of embryoid bodies collected from these cultures revealed a wide variety of differentiated cell types, including derivatives of all three embryonic germ layers. Based on their origin and demonstrated properties, these human PGC-derived cultures meet the criteria for pluripotent stem cells and most closely resemble EG cells.